Oak Ridge Documents

No. Date

43 2/18/44

Minutes of Advisory Comm. to Medical Section, Rochester
"New biological problems are presented as fission
products and 'product' become available. Info on
the biological effects of these materials is urgently
needed and these materials are now available in suitable
quantity for experimental work.
s/Stafford Warren.

4 8/29/44

Memo - Hempelmann to Oppenheimer

Summery & Side.

ساده والدوري والمعاري المانكان فالتصعير للطام

1.

2.

3. Tracer experiments on humans to determine % of Pu excreted daily. To be done at Rochester.

5 none

Memo - Stone to Compton

"Finishing Health Work of Metallurgical Project" should continue health research activities in spite of approaching end of war -- need to provide safe standards for personnel producing Pu -- to be based on experimental data.

a. Clinton Labs - (animal work) FY46

 b. UC (Hamilton) - metabolism of Pu toxicity (animal Berkeley work)

Hamilton - Berkeley (cont'd) 1/11/45 - Crocker Lab

- 4) "It is planned here at Berkeley to undertake, on a limited scale, a series of metabolic studies with Pu using human subjects"
 - 6) "I believe that a few tracer studies both with human subjects and animals...should be made"
- c. Metallurgical Lab Chicago Jacobson & Stone FY46 Studies of acute & chronic radiation effects - external and internal - Pu Bone, skin, GI tract, respiratory tract

"answers to above studies badly needed, etc."

8 3/26/45

Hempelmann to Oppenheimer

MED to help make arrangements for human tracer experiments. Suggest a hospital patient at Chicago or Rochester be chosen for injection with $1\text{--}10~\mu g$ Pu.

9 3/29/45

Oppenheimer to Warren

Enclosing above memo; endorsing requests outlined; urgency of exposure of workers makes it appropriate that MED consider medical problem more intensively than in past.

--0000682

- 2 •

	•	·
10.	4/3-4/45	Conference Summary Program to be carried out at Rochester. II(p3) Special problems Dr. Bale work to be continued in future as in past; consist almost entirely of Dr. Bale and staff, work for MED as requested, including tracer studies.
24	7/7/45	Bamilton to Stone Payment to experimental subject - to keep him in Berkeley area and continue collecting samples for analysis.
27		Friedell to Area Engineer, Berkeley Suggest purchase orders for specimens
28	8/17/45	Stone - Memo to Files Re: Warren's memo: Purpose and limitations of the Biological and Health Physics Program. p.2 - comment about moral responsibility of Univ. of Chicago. Reference to what? What is "D.S.M. Program"?
29	9/18/45	Langham to Warren Revised plan of Rochester Pu experiment sent to Bale "of course you and Friedell will have final say"
34	1/3/46	Warren Specific proposals for national distribution of radioisotopes p. 3 - statement re legal responsibilities for use of experimental material in humans. Distribution only after approval by competent medical advisors and only to M.D.'s associated with recognized medical institutions.
41	4/26/46	Friedcll to Warren Proposed program at Western Reserve University 1. Toxicity of thorium 2. Radiation tissue dosages - internally distributed

radioisotopes vs. external radiation.

•		
76	9/19/45	Ltr. Goldring to Langham Enclosing clinical history of E. C.
77		Clinical summary "Experiment I on P49&4" (')
80	10/2/45	Draft report by Goldring to Ferry, Dist. Engineer, OR "Biologic & Health Physics Section" Summary of all research activities at various labs No mention of human studies
84	4/23/46	Namilton to Howland Summary of work 15 Oct 42 to 1 April 46 & projected work to end of FY 47 Comparisons of Pu retention and excretion in rodents and man discussed (based on data from a "relatively normal human subject" (p2) - excretory record, tissue samples including bone)
108	5/14/5 7 ∻7	Memo to Files by Fidler, Chief, Tech. Info., Field Op. Policy on Medical Reports "only those experiments where actual materials were intentionally introduced into the human system need to be regarded as secret"
94	12/23/46	Draft (corrected) for item 95
95	12/24/46	Memo Nichols to Area Engineer, Berkeley Reference to progress report containing statement about intravenous injection of humans with certain radioactive substances "recommends against work on humans but also denies authority for such work under terms of Manhattan contract"
	12/24/46	Memo Brundage to Cooney (Med. Dir., Manhattan Proj., D.C.) Covering letter for Nichols memo
102	1/21/47	Ltr. from E. B. Kelley to Col. Kirkpatr ick, AEC, OR "Visit of Col. Cooney to Berkeley Area" Proposed human studies discussed with Joe Hamilton. Cooney appeared to believe these would be satisfactory to AEC but Col. made no firm statements about gov't support matter to be discussed at meeting in Washington on medical programs

CAL 3 - surviving (amputee) excreta collected at Rochester

living in Texas - billing for 2 research patient; from Rochester to ANL July 73

examined at ANL, 6/11/73, (en route to Rochester)
interview, hematology, blood chemistry, X-ray & TB count

examined by Christine Waterhouse - at SMH 6/13 to 6/26/73 summary letter 6/28/73

Letter from Brues to private physician, requesting participation in study at Rochester - 3/7/73

40-009

HP-3

Patient of Waterhouse "normal health", lives in Rochester

Hospitalized Rochester 1/28 - 2/18/73 TB measurements by Rundo (ANL) 2/8/73

Bill for research patient sent to $\ensuremath{\mathsf{ANL}}$

Memo Stehney to Rowland 1/4/73
What Waterhouse wants provided for metabolic study

40-012

HP-6 Patient of Waterhouse living in Rochester Addison's Disease

Bill for research patient to ANL

6/21 - 7/1/73 at SMH metabolic studies; blood drawn for chromosome studies $\sim 2/14/73$ at SMH

Memo Stehney to Rowland 1/4/73
What Waterhouse wants provided for metabolic study

<u>40-014</u>

HP8 Female survivor Lives in Rochester - scleraderma
"may be difficult to get for study"

L'ASL Documents

No. Date

1950 Pertaining to follow-up studies of 2 patients

X-rays "get them in a completely routine manner"
Excreta collection and analysis
Langham, Bassett, Howland, Waterhouse

Rochester Documents

<u>No.</u>	<u>Date</u>	
	1947	Correspondence pertaining to publication of scientific studies Langham, Bassett, Bale, Dowdy-Dir. Manhattan Dept. (Rochester)
4	2/18/47	Dowdy to Friedell ?inclusion of Oak Ridge Patient in write up? "It is my understanding that it was your idea and that Major Howland and Cpt. Goldring were the two who actually carried out the administration"
9	1/6/48	Memo to Bale from Dr. Blair (bracion, 5m H) Human metabolism work under Bassett to be discontinued on order from Shields Warren. Bassett to write history for files.
10	6/23/48	Transmittal of Technical Report Excretion of U (Human Studies) Several authors, including C. Waterhouse

Personal File of Stehney

No.	Date	
2	12/13/72	Stehney to Rowland - Visit P. Durbin to ANL Need for CHR money for Rochester study "Waterhouse doesn't want to upset people by telling them of their exposure" Could be handled by Rochester Human Use Committee
3	12/21/72	Rowland to Rundo "expect to hospitalize 2-3 low level Pu case in Rochester" "I intend to write to Rochester to set up this hospitalization as soon as possible"
4	12/21/72	Rowland to Shutz "outside CHR we never use the word plutonium in regard to these cases." "These individuals are of interest to us because they may have received a radioactive material at some time."
16	1/2/73	Rowland to CHR Staff Members Call Meeting on 1/3 to discuss protocol for examination of living cases in Rochester area when they enter Strong Memorial Hospital
17	1/4/73	Rowland to Waterhouse Confirm arrangements for study details "we will pay all expenses" "we can arrange payment as an inducement to participate"
21	2/5/73	Littman to Rowland Report on visit to Waterhouse "The Dean has approved participation of Dr. Forbes who will cooperate with Dr. Rundo"
30	12/14/73	Stehney to Fry Report to ANL Human Use Committee
36	3/1/74	Rowland to Waterhouse info that data will be presentedoffer to be author on paper

37		Waterhouse to Rowland Agrees with submission of abstract requests that "u. of Rochester not be implicated in this study for the time being" declines coauthorship. Alludes to discussion with Dean and Bill Neuman re AEC resolution of this matter
38	3/13/74	Rowland to Fry Supplies copies of correspondence with Waterhouse as requested.
40	4/5/74	Stehney to Sinclair Supplies copies of correspondence with Waterhouse as requested.
42	1/16/45	(p3) Project Council Meeting Dr. Stone (after presentation of i.v. Pu in dog) "Does the above data have any implications regarding the value of various types of tests which might be performed on humans?" Dr. Nickson - urine, stools, blood assays planned
45	June 46	(p2) Biochemical Survey - Monthly Report Ingestion of Pu solution by 6 people

Only No. 1 relates to human experiments with isotopes -- possibly not to Pu

15 "A report of the past, present, and future research activities for Project 48-A-1" by J. G. Hamilton 9/12/47

(Pu and several fission products)

All metabolic studies in rats, with the exception of 1 human tracer experiment, using Pu. Summary of results - no specific reference to human data.

Under "projected work, FY 1948-49" (p. 9)
Human tracer studies with certain of the fission products
and fissionable elements. Purposes: (1) to establish
metabolic behavior in man and compare to rat (using relatively
short-lived isotopes); (2) to investigate potential therapeutic
and diagnostic applications (in collaboration with Dr. R.S.
Stone at Univ. of Calif. Medical School).

18 Memo from Joe Howland to Henry A. Blair (Director, Strong Mem. Hosp.) 1/3/49

"Dr. Shields Warren's request for information concerning the clinical metabolism ward"

Tabulation of pt's studied in 1948 (names, dates, diagnosis, lab tests performed:

Only familiar name found:

2/8-2/21 Addison's Disease (no lab tests listed) (not identified - as were some patients - as "AEC interest")

Dat<u>e</u> No. 12/2/44 Memo to files - S. L. Warren 21 "Medical Experimental Program on Ra & Pu" outline of ongoing research, mostly in rats. The following data are urgently needed. c. Tracer experiments in humans -- ratios of blood level to urine & fecal excretion following intravenous injection of Pu (Bale, Hempelmann, Meeting in Chicago Dec. 4 & 5 with all involved to plan. 1/19/45 Memo to Files/Friedell 27 "Program for Product Research 5 December 44" Analytical Methods Animal Metabolism Studies & Data (rat) "In conjunction with the experimental conducted in animals, it is expected that on selected human subjects tracer studies with Pu would be made. The purpose of this would be to establish a relationship between the amount excreted and the amount retained in the body. This could serve as an excellent monitoring index to determine hazards to personnel. This is of course predicated on the idea that some conclusion as to the tolerance level of Pu in the body will be established from the experimental outlined above." Attendance - as suggested by previous memos Nos. 18 & 21 -Friedell, Bale, Cole, Warren, Hempelmann, Mastick (Los Alamos), Hamilton, Stone 41 4/4/45 Ltr. Langham to Friedell Shipment of 5 mg Pu, detailed instructions for preparing solution, injection, collection and shipment of excretaattached "Protocol of Clinical Experiment" (written after administration of Pu by Friedell) see No. 43 4/6/45 Ltr. Langham to Friedell 42 With still more explicit instructions than above 43 Ltr. Friedell to Hempelmann Encloses protocol (as in No. 41) "Everything went very smoothly"...I think we will have access to considerable clinical material here and we hope to do a number of

subjects..."

59

"Santa Fe Conversations" "JWH"

No date, but references to letters and conversations 21 May, June 2, 4 and 4-9. (Warren, Langham, Hempelmann, McKana, Friedell)

Points under debate:

- a. Number of patients on which to use produce
- b. Size of dose
- c. Significance of Pu+4 and Pu+6; soluble & insoluble forms
- d. Factors related to absorption from skin
- e. Factor about urinary excretion -- difference between urinary and stool excretion (reference to Chicago and Oak Ridge patients)
- f. Distribution of Pu in tissues of humans should be carried out in moribund patients or patients dying from malignancy particularly Hodgkin's & leukemia in young people.

Proposed experimental program

Injection of 0.06 mg/kg into 10 patients with blood and urine status normal for age.

Follow-up levels in blood, urine, stools for up to levelling off state which occurs within 8 wks-2 mo.

Biopsies of bone if possible

Tissue distribution in moribund patient Skin excretions with 238-239.

65

Detailed duty descriptions June-July 45 Ferry, Friedell, Bryan, Goldring, Howland

68 7/12/45

Memo Howland to Area Engineer, Berkeley

Re: Hamilton's concern that "experimental subject" may move out of area unless some compensation be made to him.

70,71 8/9/45

Ltrs. Stone to Friedell

Re: concern about informing public about radiation hazards to MED workers: Japanese

72

Proposed Health Program for FY 46 (outline)

for Berkeley group (Hamilton)

mentions injection experiments, including tracer studies in humans using 48

75 9/5/45

Report of meeting in Rochester (by Langham)
Detailed plan for Pu tracer study

Revised plan resulting from meeting

Outline of all details -- clinical, administrative,

0000095 scientific, etc.

No mention of informing patient

76	9/19/45	Ltr. Goldring to Langham Enclosing clinical history of E. C.
77		Clinical summary "Experiment I on P49&4" (E. Cade)
80	10/2/45	Draft report by Goldring to Ferry, Dist. Engineer, OR "Biologic & Health Physics Section" Summary of all research activities at various labs No mention of human studies
84	4/23/46	Hamilton to Howland Summary of work 15 Oct 42 to 1 April 46 & projected work to end of FY 47 Comparisons of Pu retention and excretion in rodents and man discussed (based on data from a "relatively normal human subject" (p2) - excretory record, tissue samples including bone)
108	5/14/5 7	Memo to Files by Fidler, Chief, Tech. Info., Field Op. Policy on Medical Reports "only those experiments where actual materials were intentionally introduced into the human system need to be regarded as secret"
94	12/23/46	Draft (corrected) for item 95
95	12/24/46	Memo Nichols to Area Engineer, Berkeley Reference to progress report containing statement about intravenous injection of humans with certain radioactive substances "recommends against work on humans but also denies authority for such work under terms of Manhattan contract"
	12/24/46	Memo Brundage to Cooney (Med. Dir., Manhattan Proj., D.C.) Covering letter for Nichols memo
102	1/21/47	Ltr. from E. B. Kelley to Col. Kirkpatr ick, AEC, OR "Visit of Col. Cooney to Berkeley Area" Proposed human studies discussed with Joe Hamilton. Cooney appeared to believe these would be satisfactory to AEC but Col. made no firm statements about gov't support matter to be discussed at meeting in Washington on medical programs

105 4/28/47

Ltr. from Friedell to Maj. Clarkson, AEC, OR
"don't remember man's name who received some of the
material -- there are records in the files -- "Joe
Howland actually gave the material and he might
remember, etc."

106 2/6/47

Ltr. Carroll Wilson, GM, AEC to Warren
Re Opinion of Interim Medical Advisory Committee on
Clinical Testing -- legal department to investigate
responsibility of AEC

107 4/30/47

Ltr Wilson to Warren

"All Area Managers will be notified about contents"
Opinions of Commission re approved clinical testing

- a. treatment to be given only when there is expectation that it will benefit patient
- b. decision of advisability to be made by attending physician

Same procedure to be followed as agreed upon early March-"susceptible to proof from official records that, prior to treatment, each individual patient, being in an understanding state of mind, was clearly informed of the nature of the treatment and its possible effects, and expressed his willingness to receive the treatment..."

no written release required but 2 doctors to certify in writing (and made part of official record) to patient's understanding state of mind of explanation furnished and willingness to participate.

105 57:4/+7

per p. 3